Attorney's Docket No.: 06275-233001 / Z70663-1P US

Applicant: Andrew Baxter et al.

Art Unit : Unknown

Examiner: Unknown

Serial No.:

: Herewith

Filed

Title

: NOVEL COMPOUNDS

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Commissioner for Patents Washington, D.C. 20231

PRELIMINARY AMENDMENT

Prior to examination, please amend the application as follows:

In the claims:

Cancel claims 13-19 without prejudice.

Amend claims 3-6, 9, 10, 12, 20 and 21 as follows:

-- 3. (Amended) A compound of formula (I), according to Claim 1, in which the group A is substituted as shown below in formula (Ia), where B and D are selected from CR², S, O and NR²⁵, where R² is as defined in Claim 1 and R²⁵ is hydrogen or C₁-C₆ alkyl:

$$X \longrightarrow NH_2$$
 $A \longrightarrow NH$
 $A \longrightarrow O$
 NH_2
 NH_2
 NH_2

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- 4. (Amended) A compound according to claim 1 in which the ring A is thiophene.
- 5. (Amended) A compound according to claim 1 in which R¹ represents optionally substituted phenyl.
 - 6. (Amended) A compound according to claim 1 in which R² represents H or methyl.
- 9. (Amended) A process for the preparation of a compound of formula (I), according to claim 1, which comprises:
- (a) reaction of a compound of formula (II):

$$R^2$$
 A
 O
 NH_2
 NH_2
 NH_2

wherein A, R^1 and R^2 are as defined in Claim 1 with an isocyanate (X = O) or an isothiocyanate (X = S); or

(b) reaction of compound of formula (III) with a compound of formula (IV)

wherein A, X, R¹ and R² are as defined in Claim 1 and LG represents a leaving group; or

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(c) reaction of compound of formula (V) with a compound of formula (VI)

$$R^{1}\text{-LG}$$

$$R^{2}$$

$$Metal$$

$$NH_{2}$$

$$(V)$$

$$(VI)$$

wherein A, X, R¹ and R² are as defined in Claim 1 and LG represents a leaving group;

and where necessary converting the resultant compound of formula (I), or another salt thereof, into a pharmaceutically acceptable salt thereof; or converting the resultant compound of formula (I) into a further compound of formula (I); and where desired converting the resultant compound of formula (I) into an optical isomer thereof.

- 10. (Amended) A pharmaceutical composition comprising a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in claim 1, in association with a pharmaceutically acceptable adjuvant, diluent or carrier.
- 12. (Amended) A compound of formula (I), or a pharmaceutically-acceptable salt or solvate thereof, as claimed in claim 1 for use in therapy.
- 20. (Amended) A method of treating an IKK2 mediated disease which comprises administering to a patient a therapeutically effective amount of a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in claim 1.
- 21. (Amended) A method of treating an inflammatory disease in a patient suffering from, or at risk of, said disease, which comprises administering to the patient a therapeutically effective amount of a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in claim 1,--

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REMARKS

Claims 1 to 12 and 20-25 are pending in this application, claims 13-19 having been cancelled. Claims 3-6, 9, 10, 12-15, 20 and 21 are amended to delete multiple dependency. No new matter has been added.

Attached is a marked-up version of the changes being made by the current amendment.

Applicant asks that all pending claims be examined. Please apply any charges or credits to Deposit Account No. 06-1050.

Respectfully submitted,

Date: Mre 22, 2001

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Version with markings to show changes made

In the claims:

Cancel claims 13-19 without prejudice.

Claims 3-6, 9, 10, 12, 20 and 21 have been amended as follows:

3. (Amended) A compound of formula (I), according to Claim 1 [or Claim 2], in which the group A is substituted as shown below in formula (Ia), where B and D are selected from CR2, S, O and NR²⁵, where R² is as defined in Claim 1 and R²⁵ is hydrogen or C₁-C₆ alkyl:

$$X = \begin{array}{c} NH_2 \\ NH \\ R^1 \end{array}$$
 $A = \begin{array}{c} O \\ NH_2 \end{array}$ (Ia)

- 4. (Amended) A compound according to [any one of Claims 1 to 3] claim 1 in which the ring A is thiophene.
- 5. (Amended) A compound according to [any one of Claims 1 to 4] claim 1 in which R¹ represents optionally substituted phenyl.
- 6. (Amended) A compound according to [any one of Claims 1 to 5] claim 1 in which R² represents H or methyl.
- 9. (Amended) A process for the preparation of a compound of formula (I), according to [any one of Claims 1 to 8] claim 1, which comprises:
- (a) reaction of a compound of formula (II):

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(II)

wherein A, R^1 and R^2 are as defined in Claim 1 with an isocyanate (X = O) or an isothiocyanate (X = S); or

(b) reaction of compound of formula (III) with a compound of formula (IV)

wherein A, X, R¹ and R² are as defined in Claim 1 and LG represents a leaving group; or

(c) reaction of compound of formula (V) with a compound of formula (VI)

$$R^{1}$$
-LG R^{2} A O NH_{2} NH_{2} (V) (VI)

wherein A, X, R¹ and R² are as defined in Claim 1 and LG represents a leaving group; and where necessary converting the resultant compound of formula (I), or another salt Applicant: Andrew Baxter et al. Attorney's Docket No.: 06275-233001

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thereof, into a pharmaceutically acceptable salt thereof; or converting the resultant compound of formula (I) into a further compound of formula (I); and where desired converting the resultant compound of formula (I) into an optical isomer thereof.

10. (Amended) A pharmaceutical composition comprising a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in [any one of claims 1 to 8] claim 1, in association with a pharmaceutically acceptable adjuvant, diluent or carrier.

12. (Amended) A compound of formula (I), or a pharmaceutically-acceptable salt or solvate thereof, as claimed in [any one of claims 1 to 8] <u>claim 1</u> for use in therapy.

- 20. (Amended) A method of treating an IKK2 mediated disease which comprises administering to a patient a therapeutically effective amount of a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in [any one of claims 1 to 8] claim 1.
- 21. (Amended) A method of treating an inflammatory disease in a patient suffering from, or at risk of, said disease, which comprises administering to the patient a therapeutically effective amount of a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in [any one of claims 1 to 8] claim 1.